

## SEQUENCE LISTING

### (1) GENERAL INFORMATION:

5 (i) APPLICANT: Tormo, Mar

Tari, Ana M.

Lopez-Berestein, Gabriel

10 (ii) TITLE OF INVENTION: INHIBITION OF Bcl-2 PROTEIN EXPRESSION BY  
LIPOSOMAL ANTISENSE OLIGODEOXYNUCLEOTIDES

15 (iii) NUMBER OF SEQUENCES: 7

20 (iv) CORRESPONDENCE ADDRESS:

15 (A) ADDRESSEE: Arnold, White & Durkee

(B) STREET: P.O. Box 4433

(C) CITY: Houston

(D) STATE: Texas

(E) COUNTRY: United States of America

(F) ZIP: 77210

25 (v) COMPUTER READABLE FORM:

20 (A) MEDIUM TYPE: Floppy disk

(B) COMPUTER: IBM PC compatible

25 (C) OPERATING SYSTEM: PC-DOS/MS-DOS

(D) SOFTWARE: PatentIn Release #1.0, Version #1.30

30 (vi) CURRENT APPLICATION DATA:

35 (A) APPLICATION NUMBER: US Unknown

(B) FILING DATE: Concurrently Herewith

(C) CLASSIFICATION: Unknown

40 (viii) ATTORNEY/AGENT INFORMATION:

35 (A) NAME: Wilson, Mark B.

(B) REGISTRATION NUMBER: 37,259

(C) REFERENCE/DOCKET NUMBER: UTXC:504

45 (ix) TELECOMMUNICATION INFORMATION:

(A) TELEPHONE: (512) 418-3000

(B) TELEFAX: (512) 474-7577

### (2) INFORMATION FOR SEQ ID NO:1:

50 (i) SEQUENCE CHARACTERISTICS:

45 (A) LENGTH: 18 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

55 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

CAGCGTGCGC CATCCTTC

18

(2) INFORMATION FOR SEQ ID NO:2:

- 5 (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 20 base pairs  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

ACGGTCCGCC ACTCCTTCCC

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(2) INFORMATION FOR SEQ ID NO:3:

- 20 (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 16 base pairs  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

CTGAAGGGCT TCTTCC

16

30 (2) INFORMATION FOR SEQ ID NO:4:

- 35 (i) SEQUENCE CHARACTERISTICS:  
(A) LENGTH: 5086 base pairs  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

- 40 (ix) FEATURE:  
(A) NAME/KEY: CDS  
(B) LOCATION: 1459..2175

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

GCGCCCGCCC CTCCGCGCCG CCTGCCGCC CGCCCCGCC GCTCCCGCCC GCCGCTCTCC

60

45 GTGGCCCCGC CGCGCTGCCG CCGCCGCCGC TGCCAGCGAA GGTGCCGGGG CTCCGGGCC

120

TCCCTGCCGG CGGCCGTCAG CGCTCGGAGC GAACTGCGCG ACGGGAGGTC CGGGAGGCGA

180

50 CCGTAGTCGC GCCGCCGCG AGGACCAGGA GGAGGAGAAA GGGTGCGCAG CCCGGAGGCG

240

GGGTGCGCCG GTGGGGTGCA GCGGAAGAGG GGGTCCAGGG GGGAGAACTT CGTAGCAGTC

300

ATCCTTTTA GGAAAAGAGG GAAAAAATAA AACCCCTCCCC CACCACCTCC TTCTCCCCAC

360

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	CCCTCGCCGC ACCACACACA GCGCGGGCTT CTAGCGCTCG GCACCGGCGG GCCAGGCGCG	420
	TCCTGCCTTC ATTTATCCAG CAGCTTTCG GAAAATGCAT TTGCTGTTCG GAGTTAAC	480
5	AGAAGACGAT TCCTGCCTCC GTCCCCGGCT CCTTCATCGT CCCATCTCCC CTGTCTCTCT	540
	CCTGGGGAGG CGTGAAGCGG TCCC GTGGAT AGAGATTCA GCCTGTGTCC GCGCGTGTGT	600
10	GCGCGCGTAT AAATTGCCGA GAAGGGAAA ACATCACAGG ACTTCTGCGA ATACCGGACT	660
	GAAAATTGTA ATTCACTCTGC CGCCGCCGCT GCCAAAAAAA AACTCGAGCT CTTGAGATCT	720
	CCGGTTGGGA TTCCTGCGGA TTGACATTTC TGTGAAGCAG AAGTCTGGGA ATCGATCTGG	780
15	AAATCCTCCT AATTTTACT CCCTCTCCCC CCGACTCCTG ATTCAATTGGG AAGTTCAAA	840
	TCAGCTATAA CTGGAGAGTG CTGAAGATTG ATGGGATCGT TGCCTTATGC ATTTGTTTG	900
20	GTTTACAAA AAGGAAACTT GACAGAGGAT CATGCTGTAC TTAAAAAATA CAAGTAAGTC	960
	TCGCACAGGA AATTGGTTTA ATGTAACCTT CAATGGAAAC CTTTGAGATT TTTTACTTAA	1020
	AGTGCATTG AGTAAATTAA ATTCCAGGC AGCTTAATAC ATTGTTTTA GCCGTGTAC	1080
25	TTGTAGTGTG TATGCCCTGC TTTCACTCAG TGTGTACAGG GAAACGCACC TGATTTTTA	1140
	CTTATTAGTT TGTTTTCTC TTAACCTTTC AGCATCACAG AGGAAGTAGA CTGATATTAA	1200
30	CAATACTTAC TAATAATAAC GTGCCTCATG AAATAAAGAT CCGAAAGGAA TTGGAATAAA	1260
	AATTCCCTGC GTCTCATGCC AAGAGGGAAA CACCAGAATC AAGTGTCCG CGTGATTGAA	1320
	GACACCCCT CGTCCAAGAA TGCAAAGCAC ATCCAATAAA ATAGCTGGAT TATAACTCCT	1380
35	CTTCTTCTC TGGGAAGG ATG GCG CAC GCT GGG AGA ACG GGG TAC GAC AAC	1440
	Met Ala His Ala Gly Arg Thr Gly Tyr Asp Asn	1491
	1 5 10	
40	CGG GAG ATA GTG ATG AAG TAC ATC CAT TAT AAG CTG TCG CAG AGG GGC	1539
	Arg Glu Ile Val Met Lys Tyr Ile His Tyr Lys Leu Ser Gln Arg Gly	
	15 20 25	
45	TAC GAG TGG GAT GCG GGA GAT GTG GGC GCC GCG CCC CCG GGG GCC GGC	1587
	Tyr Glu Trp Asp Ala Gly Asp Val Gly Ala Ala Pro Pro Gly Ala Ala	
	30 35 40	
50	CCC GCA CCG GGC ATC TTC TCC TCC CAG CCC GGG CAC ACG CCC CAT CCA	1635
	Pro Ala Pro Gly Ile Phe Ser Ser Gln Pro Gly His Thr Pro His Pro	
	45 50 55	

	GCC GCA TCC CGC GAC CCG GTC GCC AGG ACC TCG CCG CTG CAG ACC CCG Ala Ala Ser Arg Asp Pro Val Ala Arg Thr Ser Pro Leu Gln Thr Pro	60 65 70 75	1683
5	GCT GCC CCC GGC GCC GCG GGG CCT GCG CTC AGC CCG GTG CCA CCT Ala Ala Pro Gly Ala Ala Gly Pro Ala Leu Ser Pro Val Pro Pro	80 85 90	1731
10	GTG GTC CAC CTG GCC CTC CGC CAA GCC GGC GAC TTC TCC CGC CGC Val Val His Leu Ala Leu Arg Gln Ala Gly Asp Asp Phe Ser Arg Arg	95 100 105	1779
15	TAC CGC GGC GAC TTC GCC GAG ATG TCC AGC CAG CTG CAC CTG ACG CCC Tyr Arg Gly Asp Phe Ala Glu Met Ser Ser Gln Leu His Leu Thr Pro	110 115 120	1827
20	TTC ACC GCG CGG GGA CGC TTT GCC ACG GTG GTG GAG GAG CTC TTC AGG Phe Thr Ala Arg Gly Arg Phe Ala Thr Val Val Glu Glu Leu Phe Arg	125 130 135	1875
25	GAC GGG GTG AAC TGG GGG AGG ATT GTG GCC TTC TTT GAG TTC GGT GGG Asp Gly Val Asn Trp Gly Arg Ile Val Ala Phe Phe Glu Phe Gly Gly	140 145 150 155	1923
30	GTC ATG TGT GTG GAG AGC GTC AAC CGG GAG ATG TCG CCC CTG GTG GAC Val Met Cys Val Glu Ser Val Asn Arg Glu Met Ser Pro Leu Val Asp	160 165 170	1971
35	AAC ATC GCC CTG TGG ATG ACT GAG TAC CTG AAC CGG CAC CTG CAC ACC Asn Ile Ala Leu Trp Met Thr Glu Tyr Leu Asn Arg His Leu His Thr	175 180 185	2019
40	TGG ATC CAG GAT AAC GGA GGC TGG GAT GCC TTT GTG GAA CTG TAC GGC Trp Ile Gln Asp Asn Gly Gly Trp Asp Ala Phe Val Glu Leu Tyr Gly	190 195 200	2067
45	CCC AGC ATG CGG CCT CTG TTT GAT TTC TCC TGG CTG TCT CTG AAG ACT Pro Ser Met Arg Pro Leu Phe Asp Phe Ser Trp Leu Ser Leu Lys Thr	205 210 215	2115
50	CTG CTC AGT TTG GCC CTG GTG GGA GCT TGC ATC ACC CTG GGT GCC TAT Leu Leu Ser Leu Ala Leu Val Gly Ala Cys Ile Thr Leu Gly Ala Tyr	220 225 230 235	2163
	CTG AGC CAC AAG TGAAGTCAAC ATGCCTGCC CAAACAAATA TGCAAAAGGT Leu Ser His Lys		2215
	TCACTAAAGC AGTAGAAATA ATATGCATTG TCAGTGATGT ACCATGAAAC AAAGCTGCAG		2275
	GCTGTTAACG AAAAAATAAC ACACATATAA ACATCACACA CACAGACAGA CACACACACA		2335
	CACAACAATT AACAGTCTTC AGGCAAAACG TCGAATCAGC TATTTACTGC CAAAGGGAAA		2395

	TATCATTAT TTTTACATT ATTAAGAAAA AAGATTATT TATTAAGAC AGTCCCAC	2455
	AAACTCCGTC TTTGGAAATC CGACCCTAA TTGCCAAACA CCGCTTCGTG TGGCTCCACC	2515
5	TGGATGTTCT GTGCCTGTAA ACATAGATTG GCTTCCATG TTGTTGGCCG GATCACCATC	2575
	TGAAGAGCAG ACGGATGGAA AAAGGACCTG ATCATTGGGG AAGCTGGCTT TCTGGCTGCT	2635
10	GGAGGCTGGG GAGAAGGTGT TCATTCACTT GCATTCTTT GCCCTGGGG CGTGATATTA	2695
	ACAGAGGGAG GGTTCCCGTG GGGGAAAGTC CATGCCTCCC TGGCCTGAAG AAGAGACTCT	2755
	TTGCATATGA CTCACATGAT GCATACCTGG TGGGAGGAAA AGAGTTGGGA ACTTCAGATG	2815
15	GACCTAGTAC CCACTGAGAT TTCCACGCCG AAGGACAGCG ATGGGAAAAA TGCCCTTAAA	2875
	TCATAGGAAA GTATTTTTT AAGCTACCAA TTGTGCCGAG AAAAGCATT TAGCAATTAA	2935
20	TACAATATCA TCCAGTACCT TAAACCCTGA TTGTGTATAT TCATATATTG TGGATAACGCA	2995
	CCCCCCAAGT CCCAATACTG GCTCTGTCTG AGTAAGAAC AGAACCTCTT GGAACCTTGAG	3055
25	GAAGTGAACA TTTCGGTGAC TTCCGATCAG GAAGGCTAGA GTTACCCAGA GCATCAGGCC	3115
	GCCACAAGTG CCTGCTTTA GGAGACCGAA GTCCGCAGAA CCTACCTGTG TCCCAGCTTG	3175
	GAGGCCTGGT CCTGGAACCTG AGCCGGGCC TCACGGCCT CCTCCAGGGG TGATCAACAG	3235
30	GGTAGTGTGG TCTCCGAATG TCTGGAAGCT GATGGATGGA GCTCAGAATT CCACTGTCAA	3295
	GAAAGAGCAG TAGAGGGGTG TGGCTGGGCC TGTCACCTG GGGCCCTCCA GGTAGGCCCG	3355
	TTTCACGTG GAGCATAGGA GCCACGACCC TTCTTAAGAC ATGTATCACT GTAGAGGGAA	3415
35	GGAACAGAGG CCCTGGGCCT TCCTATCAGA AGGACATGGT GAAGGCTGGG AACGTGAGGA	3475
	GAGGCAATGG CCACGGCCA TTTGGCTGT AGCACATGGC ACGTTGGCTG TGTGCCCTTG	3535
	GCCACCTGTG AGTTAAAGC AAGGCTTAA ATGACTTGG AGAGGGTCAC AAATCCTAAA	3595
40	AGAACATTG AAGTGAGGTG TCATGGATTA ATTGACCCCT GTCTATGGAA TTACATGTAA	3655
	AACATTATCT TGTCACTGTA GTTGGTTTT ATTTGAAAAC CTGACAAAAA AAAAGTTCCA	3715
45	GGTGTGGAAT ATGGGGGTTA TCTGTACATC CTGGGGCATT AAAAAAAAT CAATGGTGGG	3775
	GAACATAAA GAAGTAACAA AAGAAGTGAC ATCTTCAGCA AATAAACTAG GAAATTTTT	3835
	TTCTTCCAG TTTAGAATCA GCCTGAAAC ATTGATGGAA TAACTCTGTG GCATTATTGC	3895
50	ATTATATACC ATTTATCTGT ATTAACCTTG GAATGTACTC TGTTCAATGT TTAATGCTGT	3955
	GGTTGATATT TCGAAAGCTG CTTTAAAAAA ATACATGCAT CTCAGCGTTT TTTGTTTT	4015

	AATTGTATTT	AGTTATGGCC	TATAACACTAT	TTGTGAGCAA	AGGTGATCGT	TTTCTGTTTG	4075	
	AGATTTTAT	CTCTTGATTC	TTCAAAAGCA	TTCTGAGAAG	GTGAGATAAG	CCCTGAGTCT	4135	
5	CAGCTACCTA	AGAAAAAACCT	GGATGTCACT	GGCCACTGAG	GAGCTTGTT	TCAACCAAGT	4195	
	CATGTGCATT	TCCACGTCAA	CAGAATTGTT	TATTGTGACA	GTTATATCTG	TTGTCCTT	4255	
10	GACCTTGT	TTGAAGGTT	TCCTCGTCCC	TGGGCAATT	CGCATTAAAT	TCATGGTATT	4315	
	CAGGATTACA	TGCATGTTG	GTTAAACCCA	TGAGATT	TCAGTTAAAAT	ATCCAGATGG	4375	
	CGAATGACCA	GCAGATTCAA	ATCTATGGT	GTTGACCTT	TAGAGAGTTG	CTTACGTGG	4435	
15	CCTGTTCAA	CACAGACCCA	CCCAGAGCCC	TCCTGCCCTC	CTTCCGCGGG	GGCTTCTCA	4495	
	TGGCTGTCCT	TCAGGGTCTT	CCTGAAATGC	AGTGGTCGTT	ACGCTCCACC	AAGAAAGCAG	4555	
	GAAACCTGTG	GTATGAAGCC	AGACCTCCCC	GGCGGGCCTC	AGGGAACAGA	ATGATCAGAC	4615	
20	CTTTGAATGA	TTCTAATTTT	TAAGCAAAAT	ATTATTTTAT	GAAAGGTTA	CATTGTCAA	4675	
	GTGATGAATA	TGGAATATCC	AATCCTGTGC	TGCTATCCTG	CCAAAATCAT	TTAATGGAG	4735	
25	TCAGTTGCA	GTATGCTCCA	CGTGGTAAGA	TCCTCCAAGC	TGCTTAGAA	GTAACAATGA	4795	
	AGAACGTGGA	CGTTTTAAT	ATAAAGCCTG	TTTGTCTT	TGTTGTTGTT	CAAACGGGAT	4855	
30	TCACAGAGTA	TTGAAAAAT	GTATATATAT	TAAGAGGTCA	CGGGGGCTAA	TTGCTAGCTG	4915	
	GCTGCCTTT	GCTGTGGGGT	TTTGTACCT	GGTTTAATA	ACAGTAAATG	TGCCAGCCT	4975	
	CTTGGCCCCA	GAACTGTACA	GTATTGTGGC	TGCAC	TTGCT	CTAAGAGTAG	TTGATGTTGC	5035
35	ATTTCCCTTA	TTGTTAAAAAA	CATGTTAGAA	GCAATGAATG	TATATAAAAG	C	5086	

(2) INFORMATION FOR SEQ ID NO:5:

- 40           (i) SEQUENCE CHARACTERISTICS:  
               (A) LENGTH: 239 amino acids  
               (B) TYPE: amino acid  
               (D) TOPOLOGY: linear

45           (ii) MOLECULE TYPE: protein

        (xii) SEQUENCE DESCRIPTION: SEQ ID NO:5:

50	Met Ala His Ala Gly Arg Thr Gly Tyr Asp Asn Arg Glu Ile Val Met
	1                        5                        10                        15
	Lys Tyr Ile His Tyr Lys Leu Ser Gln Arg Gly Tyr Glu Trp Asp Ala
	20                        25                        30

Gly Asp Val Gly Ala Ala Pro Pro Gly Ala Ala Pro Ala Pro Gly Ile  
                  35                        40                        45  
  
 Phe Ser Ser Gln Pro Gly His Thr Pro His Pro Ala Ala Ser Arg Asp  
                  50                        55                        60  
  
 Pro Val Ala Arg Thr Ser Pro Leu Gln Thr Pro Ala Ala Pro Gly Ala  
                  65                        70                        75                        80  
  
 10       Ala Ala Gly Pro Ala Leu Ser Pro Val Pro Pro Val Val His Leu Ala  
   85                        90                        95  
  
 Leu Arg Gln Ala Gly Asp Asp Phe Ser Arg Arg Tyr Arg Gly Asp Phe  
   100                        105                        110  
  
 15       Ala Glu Met Ser Ser Gln Leu His Leu Thr Pro Phe Thr Ala Arg Gly  
   115                        120                        125  
  
 Arg Phe Ala Thr Val Val Glu Glu Leu Phe Arg Asp Gly Val Asn Trp  
   130                        135                        140  
  
 20       Gly Arg Ile Val Ala Phe Phe Glu Phe Gly Gly Val Met Cys Val Glu  
   145                        150                        155                        160  
  
 25       Ser Val Asn Arg Glu Met Ser Pro Leu Val Asp Asn Ile Ala Leu Trp  
   165                        170                        175  
  
 Met Thr Glu Tyr Leu Asn Arg His Leu His Thr Trp Ile Gln Asp Asn  
   180                        185                        190  
  
 30       Gly Gly Trp Asp Ala Phe Val Glu Leu Tyr Gly Pro Ser Met Arg Pro  
   195                        200                        205  
  
 35       Leu Phe Asp Phe Ser Trp Leu Ser Leu Lys Thr Leu Leu Ser Leu Ala  
   210                        215                        220  
  
 Leu Val Gly Ala Cys Ile Thr Leu Gly Ala Tyr Leu Ser His Lys  
   225                        230                        235

40

(2) INFORMATION FOR SEQ ID NO:6:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 911 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

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- (ix) FEATURE:
  - (A) NAME/KEY: CDS
  - (B) LOCATION: 147..761

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- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

	TGATTGAAGA CACCCCCCTCG TCCAAGAACAT CAAAGCACAT CCAATAAAAT AGCTGGATTA	60
	TAACTCCTCT TCTTTCTCTG GGGGCCGTGG GGTGGGAGCT GGGGCGAGAG GTGCCGTTGG	120
5	CCCCCGTTGC TTTCCCTCTG GGAAGG ATG GCG CAC GCT GGG AGA ACG GGG TAC Met Ala His Ala Gly Arg Thr Gly Tyr	173
	1 5	
10	GAC AAC CGG GAG ATA GTG ATG AAG TAC ATC CAT TAT AAG CTG TCG CAG Asp Asn Arg Glu Ile Val Met Lys Tyr Ile His Tyr Lys Leu Ser Gln	221
	10 15 20 25	
15	AGG GGC TAC GAG TGG GAT GCG GGA GAT GTG GGC GCC GCG CCC CCG GGG Arg Gly Tyr Glu Trp Asp Ala Gly Asp Val Gly Ala Ala Pro Pro Gly	269
	30 35 40	
20	GCC GCC CCC GCA CCG GGC ATC TTC TCC TCC CAG CCC GGG CAC ACG CCC Ala Ala Pro Ala Pro Gly Ile Phe Ser Ser Gln Pro Gly His Thr Pro	317
	45 50 55	
25	CAT CCA GCC GCA TCC CGC GAC CCG GTC GCC AGG ACC TCG CCG CTG CAG His Pro Ala Ala Ser Arg Asp Pro Val Ala Arg Thr Ser Pro Leu Gln	365
	60 65 70	
30	ACC CCG GCT GCC CCC GGC GCC GCG GGG CCT GCG CTC AGC CCG GTG Thr Pro Ala Ala Pro Gly Ala Ala Gly Pro Ala Leu Ser Pro Val	413
	75 80 85	
35	CCA CCT GTG GTC CAC CTG GCC CTC CGC CAA GCC GGC GAC GAC TTC TCC Pro Pro Val Val His Leu Ala Leu Arg Gln Ala Gly Asp Asp Phe Ser	461
	90 95 100 105	
40	CGC CGC TAC CGC GGC GAC TTC GCC GAG ATG TCC AGC CAG CTG CAC CTG Arg Arg Tyr Arg Gly Asp Phe Ala Glu Met Ser Ser Gln Leu His Leu	509
	110 115 120	
45	ACG CCC TTC ACC GCG CGG GGA CGC TTT GCC ACG GTG GTG GAG GAG CTC Thr Pro Phe Thr Ala Arg Gly Arg Phe Ala Thr Val Val Glu Glu Leu	557
	125 130 135	
50	TTC AGG GAC GGG GTG AAC TGG GGG AGG ATT GTG GCC TTC TTT GAG TTC Phe Arg Asp Gly Val Asn Trp Gly Arg Ile Val Ala Phe Phe Glu Phe	605
	140 145 150	
	GGT GGG GTC ATG TGT GTG GAG AGC GTC AAC CGG GAG ATG TCG CCC CTG Gly Gly Val Met Cys Val Glu Ser Val Asn Arg Glu Met Ser Pro Leu	653
	155 160 165	
55	GTG GAC AAC ATC GCC CTG TGG ATG ACT GAG TAC CTG AAC CGG CAC CTG Val Asp Asn Ile Ala Leu Trp Met Thr Glu Tyr Leu Asn Arg His Leu	701
	170 175 180 185	



Ser Val Asn Arg Glu Met Ser Pro Leu Val Asp Asn Ile Ala Leu Trp  
165 170 175

5 Met Thr Glu Tyr Leu Asn Arg His Leu His Thr Trp Ile Gln Asp Asn  
180 185 190

Gly Gly Trp Val Gly Ala Ser Gly Asp Val Ser Leu Gly  
195 200 205